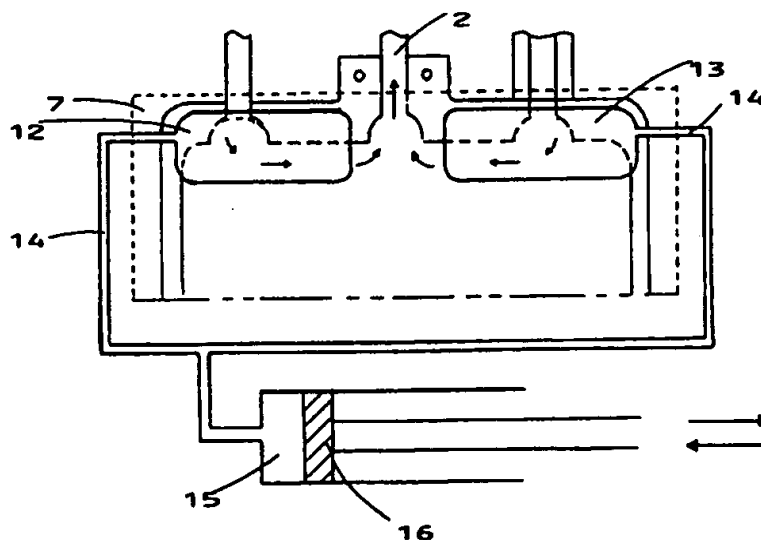




INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁶ : A61K 35/14	A1	(11) International Publication Number: WO 96/29081 (43) International Publication Date: 26 September 1996 (26.09.96)
(21) International Application Number: PCT/SE96/00356 (22) International Filing Date: 21 March 1996 (21.03.96) (30) Priority Data: 9500999-9 21 March 1995 (21.03.95) SE (71) Applicant (for all designated States except US): OMEGA MEDICINTEKNIK AB [SE/SE]; Värtavägen 13, S-115 53 Stockholm (SE). (72) Inventor; and (75) Inventor/Applicant (for US only): UNGER, Peter [SE/SE]; Värtavägen 13, S-115 53 Stockholm (SE). (74) Agent: FÖRSVARETS MATERIELVERK; Patentenheten, S- 115 88 Stockholm (SE).		(81) Designated States: AU, JP, US, European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE). Published With international search report. In English translation (filed in Swedish).

(54) Title: METHOD AND APPARATUS FOR HARVESTING BLOOD COMPONENTS



(57) Abstract

The invention relates to a method and an extractor for pressing out plasma and buffy coat from a collapsible blood container (1), in which blood has been divided, by centrifugation, into a plasma layer, a buffy coat and a layer of red blood cells. The plasma and then the buffy coat are pressed out each through a separate outlet tube (2) or through a common outlet tube (2), which is connected to the top of the blood container, a pulsating pressure being applied to a top section of the blood container (1) during the end phase of the pressing-out of buffy coat. The extractor comprises a stationary support surface (6) and a movable pressing member (7), between which the blood container is suspended and subjected to a compressive force in order to press out the plasma and then the buffy coat through the outlet tube (2). The extractor is characterised by having one or more inflatable cushions (12, 13) arranged on the support surface (6) or the pressing member (7) or both on the same level as the top section of the blood container (1), and means (15, 16) for pulsating the pressure in said cushions.

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AM	Armenia	GB	United Kingdom	MW	Malawi
AT	Austria	GE	Georgia	MX	Mexico
AU	Australia	GN	Guinea	NE	Niger
BB	Barbados	GR	Greece	NL	Netherlands
BE	Belgium	HU	Hungary	NO	Norway
BF	Burkina Faso	IE	Ireland	NZ	New Zealand
BG	Bulgaria	IT	Italy	PL	Poland
BJ	Benin	JP	Japan	PT	Portugal
BR	Brazil	KE	Kenya	RO	Romania
BY	Belarus	KG	Kyrgyzstan	RU	Russian Federation
CA	Canada	KP	Democratic People's Republic of Korea	SD	Sudan
CF	Central African Republic	KR	Republic of Korea	SE	Sweden
CG	Congo	KZ	Kazakhstan	SG	Singapore
CH	Switzerland	LI	Liechtenstein	SI	Slovenia
CI	Côte d'Ivoire	LK	Sri Lanka	SK	Slovakia
CM	Cameroon	LR	Liberia	SN	Senegal
CN	China	LT	Lithuania	SZ	Swaziland
CS	Czechoslovakia	LU	Luxembourg	TD	Chad
CZ	Czech Republic	LV	Latvia	TG	Togo
DE	Germany	MC	Monaco	TJ	Tajikistan
DK	Denmark	MD	Republic of Moldova	TT	Trinidad and Tobago
EE	Estonia	MG	Madagascar	UA	Ukraine
ES	Spain	ML	Mali	UG	Uganda
FI	Finland	MN	Mongolia	US	United States of America
FR	France	MR	Mauritania	UZ	Uzbekistan
GA	Gabon			VN	Viet Nam

Method and apparatus for harvesting blood components

The invention relates to blood component preparation. More specifically, the invention relates to a method and an extractor for pressing out plasma and buffy coat from a collapsible blood container, e.g. a blood bag, in which blood has been divided, by centrifugation, into a plasma layer, a buffy coat layer and a layer of red blood cells, the plasma and then the buffy coat being pressed out each through a separate outlet tube or through a common outlet tube, which is connected to the top of the blood container.

10 In blood component preparation, blood is donated to a blood bag via a blood draw-off tube permanently connected to the bag. Besides, a number of side bags are connected to the blood bag by means of tubes. These connections are normally collected in the upper edge of the blood bag. Subsequently, the blood bag is centrifuged upright in a centrifuge cup, such that the blood forms layers, viz. an uppermost plasma layer, a subjacent buffy coat layer whose volume is relatively small, and a lowermost layer of red blood cells. At least the plasma and buffy coat fractions are then transferred to side bags by the blood bag being suspended at its upper edge in an extractor and subjected to mechanical pressure between a support surface and a pressure plate. The plasma then flows through an outlet tube over to a plasma container, while the buffy coat slowly flows upwards in the blood container. The outlet tube is normally branched into a plasma tube extending to a plasma container and a buffy coat tube extending to a buffy coat container but, alternatively, separate outlet tubes can be provided for each container. When the buffy coat has reached the top of the blood container, the plasma tube is closed and the buffy coat tube is opened. The major part of the buffy coat now flows over to the buffy coat container. However, a certain amount of buffy coat penetrates into the indentations formed by the various connections in the upper edge of the blood container, and there is a great risk that this amount remains in the blood bag after pressing-out. The remaining amount of buffy coat will mix with the red blood cell fraction and constitute a contamination therein. In manually operated extractors, one may manually press out these residues and feed them to the outlet tube, such that they can be pressed over to the buffy coat container. However, an efficient method of reaching these buffy coat residues in automatic extractors has not been available.

An object of the present invention is to provide a method and an extractor which yield a more complete discharge of the buffy coat fraction and, thus, a purer fraction of red blood cells.

5 This is achieved by means of a method and an extractor as defined in the claims.

The method according to the invention is characterised in that a pulsating pressure is applied to a top section of the blood container during the end phase of the pressing-out of the buffy coat. At this stage of the pressing-out operation, the buffy
10 coat constitutes a narrow layer right at the very top at the edge of the blood container. The pulsating pressure gives the layer a pulsating motion and causes the buffy coat to be sucked out from said indentations in the edge of the blood container and flow freely towards the outlet tube for the buffy coat. This flow is additionally facilitated if the pulsating motion is not uniform along the entire width of the
15 blood container but is designed so as to produce a pumping effect towards the outlet tube. Such a pumping effect can be achieved, for instance, by the pulsating pressure being applied on both sides of the connection of the buffy coat outlet tube to the top of the blood container.

20 The pulsating pressure can be produced in various ways. According to a preferred embodiment, use is made of one or more inflatable cushions abutting against the top section of the blood bag, and the pressure in said cushions is caused to pulsate at a certain selected frequency.

25 The inventive extractor, which comprises a stationary support surface and a movable pressing member between which the blood container is suspended and is subjected to a compressive force, is characterised by one or more inflatable cushions arranged on the support surface or pressing member or both on the same level as the top section of the blood container, and means for pulsating the pressure in
30 said cushions.

The invention will be further described below with reference to the accompanying Figures.

35 Fig. 1 is a front projection of an embodiment of an extractor according to the invention.

Fig. 2 is a side projection of the extractor in Fig. 1.

Fig. 3 illustrates on a larger scale a top section of an extractor with two inflatable cushions and a device for pulsating the pressure therein.

5 Fig. 4 illustrates a top section of a blood container at the end of the pressing-out of the buffy coat.

Figs 1 and 2 illustrate an embodiment of an extractor consisting of a base with a support surface 6 and a pressing member 7, which is movable towards the support surface and which, in this case, consists of a pressure plate. The pressure plate is operated by a pressure cylinder 8. Between the support surface and the pressure plate, a blood container 1 is suspended at its upper edge from a suspension hook 9. From the top of the blood container extends an outlet tube 2, which in the illustrated embodiment branches into a plasma tube 10 extending to a plasma container (not shown) and a buffy coat tube 11 extending to a buffy coat container (not shown). Alternatively, the plasma container and the buffy coat container can each have an outlet tube 2 which is connected to the top of the blood container. Moreover, there are usually additional connections to the top of the blood container, for instance, a closed blood draw-off tube and a tube extending to a SAGMAN container or the like. At the top of the inside of the pressure plate, on the same level as the top section of the blood container, two inflatable cushions 12 and 13 are arranged on both sides of the connection of the outlet tube 2 to the blood container.

Fig. 3 illustrates a top section of the extractor, where the pressure plate 7 for the sake of clarity is assumed to be transparent, such that the cushions 12 and 13 and the posteriorly situated blood container 1 are clearly to be seen. The blood container is here provided with three different connections in the top, and the intermediate connection concerns a common outlet tube 2 for plasma and buffy coat. The inflatable cushions are connected via conduits 14 to a cylinder 15, which is provided with a piston 16. By a reciprocating motion of the piston, the pressure in the containers can be caused to pulsate at a selected frequency and amplitude. By exposing the cushions to a varying pressure which is higher and lower than the pressure in the blood container, the blood container part abutting against the cushions will be alternately compressed and expanded. By the pulsating volume being small relative to the volume of the blood container, its motion will be affected but to a marginal extent. The basic pressure in the blood container is determined by the pressure exerted by the pressure plate 7.

Fig. 4 illustrates the top section of the blood container during the end phase of the pressing-out of buffy coat 3 through the outlet tube 2. There are indentations 4, in which the buffy coat tends to be stuck.

5

The extractor functions in the following manner: A centrifuged blood container, in which the blood has been divided into an upper plasma layer, an intermediate buffy coat layer and a lower layer of red blood cells, is suspended from the suspension hook 9 and compressed between the support surface 6 and the pressure plate 7.

10 Plasma flows out through the outlet tube 2 and further through the plasma tube 10 to a plasma container. When the buffy coat layer has reached the upper edge of the blood container, the plasma tube 10 is closed, and instead the buffy coat tube 11 is opened. Buffy coat is pressed over to a buffy coat container and, during the final stage of this pressing-out, the pulsating of the pressure in the inflatable cushions 12 and 13 is started. The pulsating pressure applied to the top section of the
15 blood container loosens the residues of buffy coat in the indentations 4 (Fig. 4) and these can flow out through the outlet tube 2. It has been found particularly efficient, as illustrated in the Figures, to perform the pulsation on both sides of the connection of the outlet tube 2 to the blood container. A suitable frequency of the pulsation
20 can be 10-60 pulses/min. Especially good results are obtained in conventional blood bags when pulsating at a frequency of 20-40 pulses/ min.

Example

Comparative tests with and without pulsation were carried out when pressing out
25 plasma and buffy coat from a number of centrifuged blood bags. The result appears from the following Table.

Blood bag No.	Whole blood litre	Pulsation	BC litre	RBC litre	%WBC	%platelets
1	0.490	20 pulses/min	0.056	0.143	7.4	11.7
2	0.477	20 pulses/min	0.033	0.136	8.8	12.2
3	0.518	30 pulses/min	0.045	0.148	11.0	7.9
4	0.501	34 pulses/min	0.044	0.189	15.8	14.2
5	0.545	34 pulses/min	0.053	0.212	8.6	5.1
6	0.499	34 pulses/min	0.054	0.157	10.1	6.5
7	0.552	40 pulses/min	0.059	0.194	10.4	10.5
8	0.514	40 pulses/min	0.054	0.169	10.3	9.4
9	0.535	40 pulses/min	0.059	0.275	8.7	4.8
10	0.543	40 pulses/min	0.057	0.283	8.8	5.9
11	0.541	40 pulses/min	0.061	0.318	10.5	5.9
12	0.498	No	0.046	0.155	17.6	33.8
13	0.489	No	0.046	0.149	26.5	30.1
14	0.537	No	0.043	0.184	33.6	80

- 5 %WBC states the proportion of the white blood cells in the blood donation that was found in the red blood cell concentrate (RBC). % platelets states the corresponding value in respect of platelets. Thus, these values state to what extent the red blood cells were contaminated with white blood cells and platelets owing to residues of the buffy coat fraction. The Table shows that the pulsation according to the invention caused a considerably improved removal of the buffy coat fraction and, thus, a reduced contamination of the red blood cells.

Claims:

1. A method for pressing out plasma and buffy coat from a collapsible blood container (1), in which blood has been divided, by centrifugation, into a plasma layer, a buffy coat layer and a layer of red blood cells, the plasma and then the buffy coat being pressed out each through a separate outlet tube (2) or through a common outlet tube (2) which is connected to the top of the blood container, c h a r -
a c t e r i s e d in that a pulsating pressure is applied to a top section of the blood container (1) during the end phase of the pressing-out of the buffy coat.
2. The method as claimed in claim 1, c h a r a c t e r i s e d in that the pulsating pressure is applied on both sides of the connection of the buffy coat outlet tube (2) to the top of the blood container.
3. The method as claimed in claim 1 or 2, c h a r a c t e r i s e d in that the pulsating pressure is produced by means of one or more inflatable cushions (12, 13), which abut against the top section of the blood container, and that the pressure in said cushions is caused to pulsate at a certain selected frequency.
4. The method as claimed in claim 3, c h a r a c t e r i s e d in that the pressure is caused to pulsate at a frequency of 10-60 pulses/ min, preferably 20-40 pulses/min.
5. An extractor for pressing out plasma and buffy coat from a collapsible blood container (1), in which blood has been divided, by centrifugation, into a plasma layer, a buffy coat layer and a layer of red blood cells, comprising a stationary support surface (6) and a movable pressing member (7), between which the blood container is suspended and subjected to a compressive force to press out the plasma and then the buffy coat each through a separate outlet tube (2) or through a common outlet tube (2) which is connected to the top of the blood container, c h a r a c t e r i s e d by one or more inflatable cushions (12, 13) arranged on the support surface (6) or the pressing member (7) or both on the same level as the top section of the blood container (1), and means (15, 16) for pulsating the pressure in said cushions.
6. The extractor as claimed in claim 5, c h a r a c t e r i s e d in that the inflatable cushions (12, 13) are arranged on both sides of the connection of the buffy coat outlet tube (2) to the top of the blood container (1).

7. The extractor as claimed in claim 5, c h a r a c t e r i s e d in that the inflatable cushions (12, 13) communicate with a cylinder (15) having a piston (16), by means of which the pressure is pulsated.

1

INTERNATIONAL SEARCH REPORT

International application No.
PCT/SE 96/00356

A. CLASSIFICATION OF SUBJECT MATTER				
IPC6: A61K 35/14 According to International Patent Classification (IPC) or to both national classification and IPC				
B. FIELDS SEARCHED				
Minimum documentation searched (classification system followed by classification symbols)				
IPC6: A61K				
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched				
SE,DK,FI,NO classes as above				
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)				
MEDLINE, BIOSIS, EMBASE, WPI, CLAIMS, LANCET				
C. DOCUMENTS CONSIDERED TO BE RELEVANT				
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.		
A	EP 0161551 A2 (NPBI NEDERLANDS PRODUKTIELABORATORIUM VOOR BLOEDTRANSFUSIEAPPARATUUR EN INFUSIEVLOEISTOFFEN B.V.), 21 November 1985 (21.11.85), the whole document --	1-7		
A	US 5045185 A (OHNAKA ET AL), 3 Sept 1991 (03.09.91), figure 8 --	1-7		
A	US 5135646 A (TANOKURA ET AL), 4 August 1992 (04.08.92), the whole document --	1-7		
<input checked="" type="checkbox"/> Further documents are listed in the continuation of Box C. <input checked="" type="checkbox"/> See patent family annex.				
<table style="width: 100%; border: none;"> <tr> <td style="width: 50%; vertical-align: top; border: none;"> * Special categories of cited documents "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed </td> <td style="width: 50%; vertical-align: top; border: none;"> "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "Z" document member of the same patent family </td> </tr> </table>			* Special categories of cited documents "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "Z" document member of the same patent family
* Special categories of cited documents "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "Z" document member of the same patent family			
Date of the actual completion of the international search	Date of mailing of the international search report			
12 June 1996	05 -07- 1996			
Name and mailing address of the ISA/ Swedish Patent Office Box 5055, S-102 42 STOCKHOLM Facsimile No. +46 8 666 02 86	Authorized officer Ake Lindberg Telephone No. +46 8 782 25 00			

INTERNATIONAL SEARCH REPORT

International application No.

PCT/SE 96/00356

C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	US 4976851 A (TANOKURA ET AL), 11 December 1990 (11.12.90), the whole document -- -----	1-7

INTERNATIONAL SEARCH REPORT
Information on patent family members

01/04/96

International application No.
PCT/SE 96/00356

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
EP-A2- 0161551	21/11/85	SE-T3- 0161551 DE-A, A, C 3417892 JP-B- 1010229 JP-C- 1525690 JP-A- 61029363 US-A- 4663032	14/11/85 21/02/89 30/10/89 10/02/86 05/05/87
US-A- 5045185	03/09/91	AU-B, B- 611656 AU-A- 8171987 DE-A- 3783463 EP-A, A, B 0329786 SE-T3- 0329786 JP-C- 1773019 JP-B- 4065700 JP-A- 63119775 WO-A, A- 8803418 JP-C- 1810406 JP-B- 5012943 JP-A- 63119776	20/06/91 01/06/88 18/02/93 30/08/89 14/07/93 20/10/92 24/05/88 19/05/88 27/12/93 19/02/93 24/05/88
US-A- 5135646	04/08/92	AU-B, B- 621764 AU-A- 5896790 DE-D, T- 69022903 EP-A, A, B 0407999 SE-T3- 0407999 EP-A- 0593102 JP-A- 3047266	19/03/92 28/02/91 04/04/96 16/01/91 20/04/94 28/02/91
US-A- 4976851	11/12/90	AU-B, B- 619012 AU-A- 3605789 CA-A- 1331873 DE-U- 6890260 EP-A, A, B 0348682 SE-T3- 0348682 JP-A- 2098362 JP-B- 6022601	16/01/92 07/12/89 06/09/94 01/10/92 03/01/90 10/04/90 30/03/94